

SEQUENCE LISTING

<110> CARILLO, Serge
 BLANCHARD, Jean-Marie
 PIECHACZYK, Marc

<120> METHOD OF CANCER TREATMENT BY P53 PROTEIN CONTROL

<130> ST94037A-US

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<150> 08/737,953

<151> 1996-11-27

<150> FR94/06583

<151> 1994-05-31

<150> WO PCT/FR95/00670

<151> 1995-05-22

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<170> PatentIn Ver. 2.1

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<211> 2085

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gta aaa aca gaa cct gag aag aag tca cag tca acc aag ctg tct gtg 96
 Val Lys Thr Glu Pro Glu Lys Lys Ser Gln Ser Thr Lys Leu Ser Val
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gtt cat gag aaa aaa tcc caa gaa gga aag cca aaa gaa cac aca gag 144
 Val His Glu Lys Lys Ser Gln Glu Gly Lys Pro Lys Glu His Thr Glu
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cca aaa agc cta ccc aag cag gca tca gat aca gga agt aac gat gct 192
 Pro Lys Ser Leu Pro Lys Gln Ala Ser Asp Thr Gly Ser Asn Asp Ala
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cac aat aaa aaa gca gtt tcc aga tca gct gaa cag cag cca tca gag 240
 His Asn Lys Lys Ala Val Ser Arg Ser Ala Glu Gln Gln Pro Ser Glu
 65 70 75 80

aaa tca aca gaa cca aag act aaa cca caa gac atg att tct gct ggt 288
 Lys Ser Thr Glu Pro Lys Thr Lys Pro Gln Asp Met Ile Ser Ala Gly

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| | | | | | | | | | | | | | | | | | | | |
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| | | | | | 85 | | | | | 90 | | | | | 95 | | | | |
| 5 | | gga Gly | gag Glu | agt Ser | gtt Val | gct Ala | ggc Gly | atc Ile | act Thr | gca Ala | ata Ile | tct Ser | ggc Gly | aag Lys | ccg Pro | ggt Gly | gac Asp | 336 | |
| | | | | | 100 | | | | | 105 | | | | | 110 | | | | |
| 10 | | aag Lys | aaa Lys | aaa Lys | gaa Glu | aag Lys | aaa Lys | tca Ser | tta Leu | acc Thr | cca Pro | gct Ala | gtg Val | cca Pro | ggt Val | gaa Glu | tct Ser | 384 | |
| | | | | 115 | | | | | 120 | | | | | 125 | | | | | |
| | | aaa Lys | ccg Pro | gat Asp | aaa Lys | cca Pro | tcg Ser | gga Gly | aag Lys | tca Ser | ggc Gly | atg Met | gat Asp | gct Ala | gct Ala | ttg Leu | gat Asp | 432 | |
| | | | 130 | | | | | 135 | | | | | 140 | | | | | | |
| 15 | | gac Asp | tta Leu | ata Ile | gat Asp | act Thr | tta Leu | gga Gly | gga Gly | cct Pro | gaa Glu | gaa Glu | act Thr | gaa Glu | gaa Glu | gaa Glu | aat Asn | 480 | |
| | | | | | | | 150 | | | | | | 155 | | | | 160 | | |
| 20 | | aca Thr | acg Thr | tat Tyr | act Thr | gga Gly | cca Pro | gaa Glu | gtt Val | tca Ser | gat Asp | cca Pro | atg Met | agt Ser | tcc Ser | acc Thr | tac Tyr | 528 | |
| | | | | | | 165 | | | | | 170 | | | | | 175 | | | |
| 25 | | ata Ile | gag Glu | gaa Glu | ttg Leu | ggc Gly | aaa Lys | aga Arg | gaa Glu | gtc Val | aca Thr | att Ile | cct Pro | cca Pro | aaa Lys | tat Tyr | agg Arg | 576 | |
| | | | | | 180 | | | | | 185 | | | | | 190 | | | | |
| 30 | | gaa Glu | cta Leu | ttg Leu | gct Ala | aaa Lys | aag Lys | gaa Glu | ggg Gly | atc Ile | aca Thr | ggg Gly | cct Pro | cct Pro | gca Ala | gac Asp | tct Ser | 624 | |
| | | | | 195 | | | | | 200 | | | | | 205 | | | | | |
| | | tca Ser | aaa Lys | ccc Pro | ata Ile | ggg Gly | cca Pro | gat Asp | gat Asp | gct Ala | ata Ile | gac Asp | gcc Ala | ttg Leu | tca Ser | tct Ser | gac Asp | 672 | |
| | | | 210 | | | | | 215 | | | | | 220 | | | | | | |
| 35 | | ttc Phe | acc Thr | tgt Cys | ggg Gly | tcg Ser | cct Pro | aca Thr | gct Ala | gct Ala | gga Gly | aag Lys | aaa Lys | act Thr | gaa Glu | aaa Lys | gag Glu | 720 | |
| | | | | | | | 230 | | | | | 235 | | | | | 240 | | |
| 40 | | gaa Glu | tct Ser | aca Thr | gaa Glu | gtt Val | tta Leu | aaa Lys | gct Ala | cag Gln | tca Ser | gca Ala | ggg Gly | aca Thr | gtc Val | aga Arg | agt Ser | 768 | |
| | | | | | | 245 | | | | | 250 | | | | | 255 | | | |
| 45 | | gct Ala | gct Ala | cca Pro | ccc Pro | caa Gln | gag Glu | aag Lys | aaa Lys | aga Arg | aag Lys | gtg Val | gag Glu | aag Lys | gat Asp | aca Thr | atg Met | 816 | |
| | | | | | 260 | | | | | 265 | | | | | 270 | | | | |
| 50 | | agt Ser | gat Asp | caa Gln | gca Ala | ctc Leu | gag Glu | gct Ala | ctg Leu | tcg Ser | gct Ala | tca Ser | ctg Leu | ggc Gly | acc Thr | cgg Arg | caa Gln | 864 | |
| | | | | 275 | | | | | 280 | | | | | 285 | | | | | |
| | | gca Ala | gaa Glu | cct Pro | gag Glu | ctc Leu | gac Asp | ctc Leu | cgc Arg | tca Ser | att Ile | aag Lys | gaa Glu | gtc Val | gat Asp | gag Glu | gca Ala | 912 | |
| | | | | 290 | | | | 295 | | | | | 300 | | | | | | |
| 55 | | aaa Lys | gct Ala | aaa Lys | gaa Glu | gaa Glu | aaa Lys | cta Leu | gag Glu | aag Lys | tgt Cys | ggc Gly | gag Glu | gat Asp | gat Asp | gaa Glu | aca Thr | 960 | |
| | | | | | | | 310 | | | | | 315 | | | | | 320 | | |

| | | | | | | | | | | | | | | | | | | |
|----|--|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|------|
| | | atc | cca | tct | gag | tac | aga | tta | aaa | cca | gcc | acg | gat | aaa | gat | gga | aaa | 1008 |
| | | Ile | Pro | Ser | Glu | Tyr | Arg | Leu | Lys | Pro | Ala | Thr | Asp | Lys | Asp | Gly | Lys | |
| | | | | | 325 | | | | | | 330 | | | | | 335 | | |
| 5 | | cca | cta | ttg | cca | gag | cct | gaa | gaa | aaa | ccc | aag | cct | cgg | agt | gaa | tca | 1056 |
| | | Pro | Leu | Leu | Pro | Glu | Pro | Glu | Glu | Lys | Pro | Lys | Pro | Arg | Ser | Glu | Ser | |
| | | | | | 340 | | | | | 345 | | | | | 350 | | | |
| 10 | | gaa | ctc | att | gat | gaa | ctt | tca | gaa | gat | ttt | gac | cgg | tct | gaa | tgt | aaa | 1104 |
| | | Glu | Leu | Ile | Asp | Glu | Leu | Ser | Glu | Asp | Phe | Asp | Arg | Ser | Glu | Cys | Lys | |
| | | | | 355 | | | | | 360 | | | | | 365 | | | | |
| | | gag | aaa | cca | tct | aag | cca | act | gaa | aag | aca | gaa | gaa | tct | aag | gcc | gct | 1152 |
| 15 | | Glu | Lys | Pro | Ser | Lys | Pro | Thr | Glu | Lys | Thr | Glu | Glu | Ser | Lys | Ala | Ala | |
| | | | 370 | | | | | 375 | | | | | 380 | | | | | |
| | | gct | cca | gct | cct | gtg | tcg | gag | gct | gtg | tct | cgg | acc | tcc | atg | tgt | agt | 1200 |
| | | Ala | Pro | Ala | Pro | Val | Ser | Glu | Ala | Val | Ser | Arg | Thr | Ser | Met | Cys | Ser | |
| | | 385 | | | | | 390 | | | | | 395 | | | | | 400 | |
| 20 | | ata | cag | tca | gca | ccc | cct | gag | ccg | gct | acc | ttg | aag | ggc | aca | gtg | cca | 1248 |
| | | Ile | Gln | Ser | Ala | Pro | Pro | Glu | Pro | Ala | Thr | Leu | Lys | Gly | Thr | Val | Pro | |
| | | | | | 405 | | | | | | 410 | | | | | 415 | | |
| 25 | | gat | gat | gct | gta | gaa | gcc | ttg | gct | gat | agc | ctg | ggg | aaa | aag | gaa | gca | 1296 |
| | | Asp | Asp | Ala | Val | Glu | Ala | Leu | Ala | Asp | Ser | Leu | Gly | Lys | Lys | Glu | Ala | |
| | | | | | 420 | | | | 425 | | | | | 430 | | | | |
| 30 | | gat | cca | gaa | gat | gga | aaa | cct | gtg | atg | gat | aaa | gtc | aag | gag | aag | gcc | 1344 |
| | | Asp | Pro | Glu | Asp | Gly | Lys | Pro | Val | Met | Asp | Lys | Val | Lys | Glu | Lys | Ala | |
| | | | | 435 | | | | | 440 | | | | | 445 | | | | |
| | | aaa | gaa | gaa | gac | cgt | gaa | aag | ctt | ggt | gaa | aaa | gaa | gaa | aca | att | cct | 1392 |
| 35 | | Lys | Glu | Glu | Asp | Arg | Glu | Lys | Leu | Gly | Glu | Lys | Glu | Glu | Thr | Ile | Pro | |
| | | | 450 | | | | | 455 | | | | | 460 | | | | | |
| | | cct | gat | tat | aga | tta | gaa | gag | gtc | aag | gat | aaa | gat | gga | aag | cca | ctc | 1440 |
| | | Pro | Asp | Tyr | Arg | Leu | Glu | Glu | Val | Lys | Asp | Lys | Asp | Gly | Lys | Pro | Leu | |
| | | 465 | | | | | 470 | | | | | 475 | | | | | 480 | |
| 40 | | ctg | cca | aaa | gag | tct | aag | gaa | cag | ctt | cca | ccc | atg | agt | gaa | gac | ttc | 1488 |
| | | Leu | Pro | Lys | Glu | Ser | Lys | Glu | Gln | Leu | Pro | Pro | Met | Ser | Glu | Asp | Phe | |
| | | | | | 485 | | | | | | 490 | | | | | 495 | | |
| 45 | | ctt | ctg | gat | gct | ttg | tct | gag | gac | ttc | tct | ggt | cca | caa | aat | gct | tca | 1536 |
| | | Leu | Leu | Asp | Ala | Leu | Ser | Glu | Asp | Phe | Ser | Gly | Pro | Gln | Asn | Ala | Ser | |
| | | | | | 500 | | | | | 505 | | | | | 510 | | | |
| 50 | | tct | ctt | aaa | ttt | gaa | gat | gct | aaa | ctt | | | | | | | | |

| | 545 | | 550 | | 555 | | 560 | |
|----|---|-----|-----|--|-----|--|-----|------|
| 5 | tct gac agt cta gga caa agg cag cct gac cca gat gag aac aaa cca | | | | | | | 1728 |
| | Ser Asp Ser Leu Gly Gln Arg Gln Pro Asp Pro Asp Glu Asn Lys Pro | | | | | | | |
| | | 565 | | | 570 | | 575 | |
| 10 | atg gga gat aaa gta aag gaa aaa gct aaa gct gaa cat aga gac aag | | | | | | | 1776 |
| | Met Gly Asp Lys Val Lys Glu Lys Ala Lys Ala Glu His Arg Asp Lys | | | | | | | |
| | | 580 | | | 585 | | 590 | |
| 15 | ctt gga gaa aga gat gac act atc cca cct gaa tac aga cat ctc ctg | | | | | | | 1824 |
| | Leu Gly Glu Arg Asp Asp Thr Ile Pro Pro Glu Tyr Arg His Leu Leu | | | | | | | |
| | | 595 | | | 600 | | 605 | |
| 20 | gat gat aat gga cag gac aaa cca gtg aag cca cct aca aag aaa tca | | | | | | | 1872 |
| | Asp Asp Asn Gly Gln Asp Lys Pro Val Lys Pro Pro Thr Lys Lys Ser | | | | | | | |
| | | 610 | | | 615 | | 620 | |
| 25 | gag gat tca aag aaa cct gca gat gac caa gac ccc att gat gct ctc | | | | | | | 1920 |
| | Glu Asp Ser Lys Lys Pro Ala Asp Asp Gln Asp Pro Ile Asp Ala Leu | | | | | | | |
| | | 625 | | | 630 | | 635 | 640 |
| 30 | tca gga gat ctg gac agc tgt ccc tcc act aca gaa acc tca cag aac | | | | | | | 1968 |
| | Ser Gly Asp Leu Asp Ser Cys Pro Ser Thr Thr Glu Thr Ser Gln Asn | | | | | | | |
| | | 645 | | | 650 | | 655 | |
| 35 | aca gca aag gat aag tgc aag aag gct gct tcc agc tcc aaa gca cct | | | | | | | 2016 |
| | Thr Ala Lys Asp Lys Cys Lys Lys Ala Ala Ser Ser Ser Lys Ala Pro | | | | | | | |
| | | 660 | | | 665 | | 670 | |
| 40 | aag aat gga ggt aaa gcg aag gat tca gca aag aca aca gag gaa act | | | | | | | 2064 |
| | Lys Asn Gly Gly Lys Ala Lys Asp Ser Ala Lys Thr Thr Glu Glu Thr | | | | | | | |
| | | 675 | | | 680 | | 685 | |
| 45 | tcc aag cca aaa gat gac taa | | | | | | | 2085 |
| | Ser Lys Pro Lys Asp Asp | | | | | | | |
| | | 690 | | | 695 | | | |
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| | <212> PRT | | | | | | | |
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| 55 | <400> 2 | | | | | | | |
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| | 1 5 10 15 | | | | | | | |
| | Val Lys Thr Glu Pro Glu Lys Lys Ser Gln Ser Thr Lys Leu Ser Val | | | | | | | |
| | 20 25 30 | | | | | | | |
| 60 | Val His Glu Lys Lys Ser Gln Glu Gly Lys Pro Lys Glu His Thr Glu | | | | | | | |
| | 35 40 45 | | | | | | | |
| | Pro Lys Ser Leu Pro Lys Gln Ala Ser Asp Thr Gly Ser Asn Asp Ala | | | | | | | |
| | 50 55 60 | | | | | | | |
| 65 | His Asn Lys Lys Ala Val Ser Arg Ser Ala Glu Gln Gln Pro Ser Glu | | | | | | | |
| | 65 70 75 80 | | | | | | | |
| 70 | Lys Ser Thr Glu Pro Lys Thr Lys Pro Gln Asp Met Ile Ser Ala Gly | | | | | | | |
| | 85 90 95 | | | | | | | |
| 75 | Gly Glu Ser Val Ala Gly Ile Thr Ala Ile Ser Gly Lys Pro Gly Asp | | | | | | | |

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| | | | | | | | | | | | | | | | | |
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| | | | | 100 | | | | | 105 | | | | | 110 | | |
| | Lys | Lys | Lys | Glu | Lys | Lys | Ser | Leu | Thr | Pro | Ala | Val | Pro | Val | Glu | Ser |
| | | | 115 | | | | | 120 | | | | | 125 | | | |
| 5 | Lys | Pro | Asp | Lys | Pro | Ser | Gly | Lys | Ser | Gly | Met | Asp | Ala | Ala | Leu | Asp |
| | | 130 | | | | | 135 | | | | | 140 | | | | |
| | Asp | Leu | Ile | Asp | Thr | Leu | Gly | Gly | Pro | Glu | Glu | Thr | Glu | Glu | Glu | Asn |
| | 145 | | | | | 150 | | | | | 155 | | | | | 160 |
| | Thr | Thr | Tyr | Thr | Gly | Pro | Glu | Val | Ser | Asp | Pro | Met | Ser | Ser | Thr | Tyr |
| | | | | | 165 | | | | | 170 | | | | | 175 | |
| 10 | Ile | Glu | Glu | Leu | Gly | Lys | Arg | Glu | Val | Thr | Ile | Pro | Pro | Lys | Tyr | Arg |
| | | | | 180 | | | | | 185 | | | | | 190 | | |
| | Glu | Leu | Leu | Ala | Lys | Lys | Glu | Gly | Ile | Thr | Gly | Pro | Pro | Ala | Asp | Ser |
| | | | 195 | | | | | 200 | | | | | 205 | | | |
| | Ser | Lys | Pro | Ile | Gly | Pro | Asp | Asp | Ala | Ile | Asp | Ala | Leu | Ser | Ser | Asp |
| 15 | | 210 | | | | | 215 | | | | | 220 | | | | |
| | Phe | Thr | Cys | Gly | Ser | Pro | Thr | Ala | Ala | Gly | Lys | Lys | Thr | Glu | Lys | Glu |
| | 225 | | | | | 230 | | | | | 235 | | | | | 240 |
| | Glu | Ser | Thr | Glu | Val | Leu | Lys | Ala | Gln | Ser | Ala | Gly | Thr | Val | Arg | Ser |
| | | | | | 245 | | | | | 250 | | | | | 255 | |
| 20 | Ala | Ala | Pro | Pro | Gln | Glu | Lys | Lys | Arg | Lys | Val | Glu | Lys | Asp | Thr | Met |
| | | | | 260 | | | | | 265 | | | | | 270 | | |
| | Ser | Asp | Gln | Ala | Leu | Glu | Ala | Leu | Ser | Ala | Ser | Leu | Gly | Thr | Arg | Gln |
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| | Ala | Glu | Pro | Glu | Leu | Asp | Leu | Arg | Ser | Ile | Lys | Glu | Val | Asp | Glu | Ala |
| 25 | | 290 | | | | | 295 | | | | | 300 | | | | |
| | Lys | Ala | Lys | Glu | Glu | Lys | Leu | Glu | Lys | Cys | Gly | Glu | Asp | Asp | Glu | Thr |
| | 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| | Ile | Pro | Ser | Glu | Tyr | Arg | Leu | Lys | Pro | Ala | Thr | Asp | Lys | Asp | Gly | Lys |
| | | | | | 325 | | | | | 330 | | | | | 335 | |
| 30 | Pro | Leu | Leu | Pro | Glu | Pro | Glu | Glu | Lys | Pro | Lys | Pro | Arg | Ser | Glu | Ser |
| | | | | 340 | | | | | 345 | | | | | 350 | | |
| | Glu | Leu | Ile | Asp | Glu | Leu | Ser | Glu | Asp | Phe | Asp | Arg | Ser | Glu | Cys | Lys |
| | | | 355 | | | | | 360 | | | | | 365 | | | |
| | Glu | Lys | Pro | Ser | Lys | Pro | Thr | Glu | Lys | Thr | Glu | Glu | Ser | Lys | Ala | Ala |
| 35 | | 370 | | | | | 375 | | | | | 380 | | | | |
| | Ala | Pro | Ala | Pro | Val | Ser | Glu | Ala | Val | Ser | Arg | Thr | Ser | Met | Cys | Ser |
| | 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| | Ile | Gln | Ser | Ala | Pro | Pro | Glu | Pro | Ala | Thr | Leu | Lys | Gly | Thr | Val | Pro |
| | | | | | 405 | | | | | 410 | | | | | 415 | |
| 40 | Asp | Asp | Ala | Val | Glu | Ala | Leu | Ala | Asp | Ser | Leu | Gly | Lys | Lys | Glu | Ala |
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| | Asp | Pro | Glu | Asp | Gly | Lys | Pro | Val | Met | Asp | Lys | Val | Lys | Glu | Lys | Ala |
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565 570 575
Met Gly Asp Lys Val Lys Glu Lys Ala Lys Ala Glu His Arg Asp Lys
580 585 590
5 Leu Gly Glu Arg Asp Asp Thr Ile Pro Pro Glu Tyr Arg His Leu Leu
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Asp Asp Asn Gly Gln Asp Lys Pro Val Lys Pro Pro Thr Lys Lys Ser
610 615 620
Glu Asp Ser Lys Lys Pro Ala Asp Asp Gln Asp Pro Ile Asp Ala Leu
625 630 635 640
10 Ser Gly Asp Leu Asp Ser Cys Pro Ser Thr Thr Glu Thr Ser Gln Asn
645 650 655
Thr Ala Lys Asp Lys Cys Lys Lys Ala Ala Ser Ser Ser Lys Ala Pro
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Ser Asp Pro Met Ser Ser Thr Tyr Ile Glu Glu Leu Gly Lys Arg Glu
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Val Thr Ile Pro Pro Lys Tyr Arg Glu Leu Leu Ala Lys Lys Glu Gly
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Ile Thr Gly Pro Pro Ala Asp Ser Ser Lys Pro Ile Gly Pro Asp Asp
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Ala Ile Asp Ala Leu Ser Ser Asp Phe Thr Cys Gly Ser Pro Thr Ala
85 90 95

gct gga aag aaa act gaa aaa gag gaa tct aca gaa gtt tta aaa gct 336
Ala Gly Lys Lys Thr Glu Lys Glu Glu Ser Thr Glu Val Leu Lys Ala
100 105 110

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